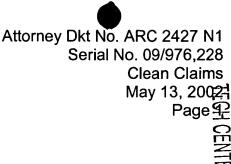
Page 42





What is claimed is:

- 1. A rate controlling membrane for an implantable drug delivery device characterized by being subjected to an elevated temperature of about 30° C to about 5° C below the melting temperature of the membrane polymer for a predetermined period of about 1 - 250 hours and subsequently incorporated into the delivery device.
- 3. A rate controlling membrane according to claim 1 wherein the membrane comprises polyurethanes or polyether blocked amides copolymers.
- 10. A rate controlling membrane according to claim 3 wherein the membrane comprises polyurethanes
- 11. A rate controlling membrane according to claim 1wherein the membrane is positioned in sealing relationship with an internal surface of one end of an impermeable reservoir of a fluid-imbibing drug delivery device, wherein said fluid imbibing drug delivery device comprises an impermeable reservoir containing a piston that divides the reservoir into a drug containing chamber and a water-swellable agent containing chamber, wherein the water-swellable agent containing chamber is provided with an outlet which accommodates said membrane.
- 12. A rate controlling membrane according to claim 3 wherein the drug containing chamber comprises leuprolide.
- 13. A rate controlling membrane according to claim 1 wherein the elevated temperature is about 45 - 80° C and the predetermined period is about 1-75 hours.
- 14. A rate controlling membrane for an implantable drug delivery device characterized by being subjected to an elevated temperature of about 30° C to about 5° C below the melting temperature of the membrane polymer for a predetermined period of about 1 to 250 hours and subsequently incorporated into

the delivery device wherein the membrane is cooled to ambient conditions before being incorporated into the delivery device.

- 15. A rate controlling membrane according to claim 3 wherein the elevated temperature is about 52 72° C and the predetermined time is about 2 36 hours.
- 16. A rate controlling membrane according to claim 10 wherein the elevated temperature is about 55 75° C and the predetermined time is about 12 48 hours.
- 17. A method for processing rate controlling membranes used in implantable drug delivery devices comprising:
- a) allowing the membrane to relax at room temperature for about12 hours to 7 days before being subjected to elevated temperature;
- b) exposing the membrane to a predetermined temperature of from about 30° C to about 5°C below the melting temperature of the membrane polymer;
- c) maintaining the membrane at the predetermined temperature for a period of time of from about 1 to 250 hours; and
- d) incorporating said membrane into a controlled drug delivery device.
- 18. A method according to claim 17 wherein the predetermined temperature is from about 45° C to 80° C.
- 19. A method according to claim 18 wherein the membrane is maintained at the predetermined temperature for a period of time of from about 1 to 75 hours.
- 20. A method according to claim 17 wherein the membrane is cooled to ambient conditions over a period of time of about 0.1-150 hours prior to incorporating the membrane into the device.

- 22. A method according to claim 17 wherein the membrane is formed from a material selected from the group consisting of polyurethanes and polyether blocked amides copolymers.
- 28. A method according to claim 17 wherein the membrane is allowed to set at ambient conditions for a period of at least about 12 hours after processing prior to exposing the membrane to said predetermined temperature.
- 29. A method according to claim 28 wherein the membrane is allowed to set at ambient conditions for a period of at least 48 hours after processing prior to exposing the membrane to said predetermined temperature.
- 30. A method according to claim 17 wherein the membrane comprises polyurethane.
- 31. A method according to claim 30 wherein the predetermined temperature is about 55 75° C and the period of time is about 12 48 hours.
- 32. A method according to claim 31 wherein the membrane is positioned in sealing relationship with an internal surface of one end of an impermeable reservoir of a fluid-imbibing drug delivery device, wherein said fluid imbibing drug delivery device comprises an impermeable reservoir containing a piston that divides the reservoir into an active agent containing chamber and a water-swellable agent containing chamber, wherein the water-swellable agent containing chamber is provided with an outlet which accommodates said membrane.
- 33. A method according to claim 32 wherein the membrane is plugshaped.
- 34. A rate controlling membrane according to claim 1 wherein the membrane comprises polyether blocked amides copolymers.
- 35. A rate controlling membrane according to claim 10 wherein the polyurethane is a single aliphatic polyether polyurethane or a blend of aliphatic polyether polyurethanes.

- 36. A rate controlling membrane according to claim 11 wherein the drug containing chamber comprises an opioid analgesic drug.
- 37. A rate controlling membrane according to claim 11 wherein the drug containing chamber comprises an antiviral drug.
- 38. A rate controlling membrane according to claim 11 wherein the drug containing chamber comprises an antineoplastic drug.
- 39. A rate controlling membrane according to claim 10 wherein the membrane is allowed to relax at room temperature for about 12 hours to 7 days before being annealed.
- 40. A rate controlling membrane for an implantable drug delivery device characterized by being subjected to an elevated temperature of about 30° C to about 5° C below the melting temperature of the membrane polymer for a predetermined period of about 1 to 250 hours and subsequently incorporated into the delivery device wherein the membrane is allowed to relax at room temperature for about 12 hours to 7 days before being annealed.
- 41 A rate controlling membrane for an implantable drug delivery device characterized by being subjected to an elevated temperature of about 52° C to about 72° C for a predetermined period of about 2 to 36 hours and subsequently incorporated into the delivery device.
- 42. A rate controlling membrane for an implantable drug delivery device characterized by being subjected to an elevated temperature of about 52° C to about 72° C for a predetermined period of about 2 to 36 hours and subsequently incorporated into the delivery device wherein the membrane is cooled to ambient conditions before being incorporated into the delivery device.
- 43. A rate controlling membrane for an implantable drug delivery device characterized by being subjected to an elevated temperature of about 52° C to about 72° C for a predetermined period of about 2 to 36 hours and subsequently incorporated into the delivery device, wherein the membrane is allowed to relax

at room temperature for about 12 hours to 7 days before being subjected to an elevated temperature.

- 44. A rate controlling membrane for an implantable drug delivery device characterized by being subjected to an elevated temperature of about 52° C to about 72° C for a predetermined period of about 2 to 36 hours and subsequently incorporated into the delivery device wherein during processing the membrane is dried to about 0 to about 1 % moisture content before being annealed and wherein the membrane is kept at about 0 to about 1% moisture content during annealing.
- 45. A rate controlling membrane for an implantable drug delivery device characterized by allowing the membrane to relax at room temperature for about 12 hours to 7 days before being annealed; subjecting the membrane to an elevated temperature of about 52° C to about 72° C for a predetermined period of about 2 to 36 hours; and cooling the membrane to ambient conditions before being incorporated into the delivery device.
- 46. A rate controlling membrane for an implantable drug delivery device characterized by allowing the membrane to relax at room temperature for about 12 hours to 7 days before being annealed; drying the membrane to about 1 to 2% moisture content; subjecting the membrane to an elevated temperature of about 52° C to about 72° C for a predetermined period of about 2 to 36 hours while keeping the moisture content of the membrane at about 1 to 2%; and cooling the membrane to ambient conditions before being incorporated into the delivery device.
- 47. A rate controlling membrane according to claim 10 wherein the elevated temperature is about 50 80° C and the predetermined time is about 4 hours 72 hours.
- 48. A method for processing rate controlling membranes used in implantable drug delivery devices comprising:

- a) allowing the membrane to relax at room temperature for about12 hours to 7 days;
- b) exposing the relaxed membrane to a predetermined temperature of from about 30° C to about 5°C below the melting temperature of the membrane polymer;
- c) maintaining the membrane at the predetermined temperature for a period of time of from about 1 to 250 hours; and
- d) incorporating said membrane into a controlled drug delivery device.
- 49. A method for processing rate controlling membranes used in implantable drug delivery devices comprising:
- a) allowing the membrane to relax at room temperature for about
 12 hours 7 days;
- b) exposing the relaxed membrane to a predetermined temperature of from about 30° C to about 5°C below the melting temperature of the membrane polymer;
- c) maintaining the membrane at the predetermined temperature for a period of time of from about 1 to 250 hours;
- d) allowing the annealed membrane to cool to room temperature for about 0.1 to 250 hours; and
- e) incorporating said membrane into a controlled drug delivery device.
- 50. A method according to claim 17 wherein the membrane comprises polyether blocked amides copolymers.
- 51 A method according to claim 50 wherein the predetermined temperature is about 55-75° C and the period of time is about 12 48 hours.
- 52 A method according to claim 51 wherein the membrane is positioned in sealing relationship with an internal surface of one end of an

impermeable reservoir of a fluid-imbibing drug delivery device, wherein said fluid imbibing drug delivery device comprises an impermeable reservoir containing a piston that divides the reservoir into an active agent containing chamber and a water-swellable agent containing chamber, wherein the water-swellable agent containing chamber is provided with an outlet which accommodates said membrane.

- 53. A rate controlling membrane for an implantable drug delivery device characterized by being subjected to an elevated temperature of about 45° C to about 80° C for a predetermined period of about 1 75 hours and subsequently incorporated into the delivery device.
- 54. A method for processing rate controlling membranes with low variability of water uptake from membrane to membrane for an implantable drug delivery device comprising:
- a) allowing the membrane to relax at room temperature for about 12 hours7 days;
 - b) drying the moisture content of the membrane to about 0 to about 1%;
- c) exposing the relaxed membrane to a predetermined temperature of from about 30° C to about 5°C below the melting temperature of the membrane polymer while maintaining the low moisture content of the membrane;
- d) maintaining the membrane at the predetermined temperature for a period of time of from about 1 to 250 hours;
- e) allowing the annealed membrane to cool to room temperature for about 0.1 to 250 hours; and
 - f) incorporating said membrane into a controlled drug delivery device.
- 55. A rate controlling membrane for an implantable drug delivery device with decreased variability of water uptake from membrane to membrane.
- 56. A rate controlling membrane for an implantable drug delivery device characterized by being subjected to an elevated temperature of about 55° C 75°

C for a predetermined period of about 12 – 48 hours wherein the membrane comprises a material selected from the group consisting of polyurethanes or polyether blocked amides copolymers.

- 57. A method for processing rate controlling membranes used in implantable drug delivery devices comprising:
- a) allowing the membrane to relax at room temperature for about12 hours to 7 days before being subjected to elevated temperature;
- b) exposing the membrane to a predetermined temperature of from about 45° C to about 80°C;
- c) maintaining the membrane at the predetermined temperature for a period of time of from about 1 to 250 hours; and
- d) incorporating said membrane into a controlled drug delivery device.
- 58. A method for processing rate controlling membranes used in implantable drug delivery devices comprising:
- a) allowing the membrane to relax at room temperature for about12 hours to 7 days before being subjected to elevated temperature;
- a) exposing the membrane to a predetermined temperature of from about 45° C to about 80°C;
- b) maintaining the membrane at the predetermined temperature for a period of time of from about 1 to 75 hours; and
- c) incorporating said membrane into a controlled drug delivery device.
- 59. A rate controlling membrane according to claim 3 wherein the membrane comprises polyether blocked amides copolymers.
- 60. An annealed rate controlling membrane for an implantable drug delivery device wherein the annealed membrane exhibits more stable water uptake and more stable water permeability than a non-annealed membrane.

- 61. An annealed rate controlling membrane for an implantable drug delivery device wherein the annealing process decreases the variability of water uptake from membrane to membrane over time.
- 62. A rate controlling membrane according to claim 1 wherein the drug containing chamber comprises leuprolide.
- 63. A rate controlling membrane according to claim 10 wherein the drug containing chamber comprises leuprolide.